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20: Boyer S, David JP, Rey D, Lemperiere G, Ravanel P.

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=> s megalosamine biosynthetic gene cluster  
L1 0 MEGALOSAMINE BIOSYNTHETIC GENE CLUSTER

=> s megosamine  
L2 52 MEGOSAMINE

$\equiv$  > s 12 and (megK)

ES Z, EZ AND (MEGR)

L3 ANSWER 1 OF 27 BIOTEC  
TI Novel isolated purified

polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide; involving vector-mediated gene transfer and expression in host cell for polyketide production

AN 2004-10434 BIOTECHDS

AB DERWENT ABSTRACT:  
NOVELTY - An isolated, purified, or recombinant nucleic acid (I), comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following: (1) an isolated, purified, or recombinant nucleic acid (II) comprising genes for the biosynthesis mycarose for attachment to a polyketide, the enzymes comprising the MegM, MegL, MegBIII, MegBIV, MegDIV, MegBII-2, and MegBVI enzymes; (2) an isolated, purified, or recombinant nucleic acid (III) comprising genes for the biosynthesis mycarose for attachment of **megosamine** of a polyketide, the enzymes comprising the MegM, MegL, MegCII, MegBVI, MegDIV, MegDV, MegDII, MegDIII, and MegDI enzymes; (3) an isolated, purified, or recombinant nucleic acid (IV) comprising genes for the biosynthesis of desosamine to a polyketide, the enzymes consisting of the MegM, MegL, MegCII, MegCIV, MegCV, MegDII, and MegDIII enzymes; (4) an expression vector (V) comprising (I); (5) a host cell comprising (I); (6) a host cell comprising (II) that expresses a polyketide modifying enzyme encoded by a gene from a mycarose biosynthetic gene set, where the enzyme is chosen from MegM, MegL, MegBIII, MegBIV, MegDIV, MegBII-2, and MegBVI, MegBV, and MegF; (7) a host cell comprising (III) that expresses a polyketide modifying enzyme encoded by a gene from a **megosamine** biosynthetic gene set, where the enzyme is chosen from MegM, MegL, MegCII, MegCIV, MegCV, MegDII, and MegDIII; and (8) a host cell comprising (IV) that expresses a polyketide modifying enzyme encoded by a gene from a desosamine biosynthetic gene set, where the enzyme is chosen from MegM, MegL, MegCII, MegCIV, MegCV, MegDII, and MegDIII, and MegCIII.

BIOTECHNOLOGY - Preferred Nucleic Acid: In (I), the gene encodes a polyketide modifying enzymes chosen from MegR, **MegK**, MegCV, MegCIV, MegBVI, MegF, MegBII, MegM, and MegL. (I) further comprises gene encoding an enzyme for the attachment of mycarose to the polyketide, preferably MegBV enzyme. (I) further comprises gene encoding an enzyme for hydroxylation of the polyketide, preferably MegF enzyme. (IV) further comprises gene encoding an enzyme for the attachment of desosamine to the polyketide, preferably MegCIII enzyme. The polyketide modifying gene is operably linked to heterologous promoter.

USE - (M1) is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising (I) under conditions in which the cell expresses a product of a gene encoded by (I) under conditions in which the unmodified polyketide is present, and producing the modified polyketide. In (M1), the cell further comprises (I) one or more module of a polyketide synthase. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. (All claimed.)

EXAMPLE - Isolation of the megalomicin biosynthetic gene cluster was as follows. A cosmid library was prepared in SuperCos vectors from Micromonospora megalomicea total DNA partially digested with Sau3AI and introduced into Escherichia coli using a Gigapack III XL in vitro packaging kit. 32P-labeled DNA probes encompassing the KS2 domain from DEBS, or a mixture of segments encompassing modules 1 and 2 from DEBS, were used separately to screen the cosmid library by colony hybridization. Several colonies which hybridized with the probes were further analyzed by sequencing the ends of their cosmid inserts using T3 and T7 primers. BLAST analysis of the sequences revealed several colonies with DNA sequences highly homologous to genes from the ery cluster. Together with restriction analysis, that led to the isolation of two overlapping cosmids, pKOSO79-93A and pKOSO79-93D which covered 45 kb of the meg cluster. A 400 base pair PCR fragment was generated from the left end of pKOSO79-93D and used to reprobe the cosmid library. Likewise, a 200 base pair PCR fragment generated from the right end of pKOSO79-93A was used to reprobe the cosmid library. Analysis of hybridizing colonies, as described above, resulted in identification of two additional cosmids pKOSO79-938B adjacent to the 5' end of pKOSO79-93D and pKOSO05.57-2.3B which overlapped the 3' ends of pKOSO79-93A and pKOSO79-93D cosmids. BLAST analysis of the far left and right end sequences of these cosmids

indicated no homology to any known genes related to polyketide biosynthesis, and therefore indicates that the set of four cosmids spans the entire megalomicin biosynthetic gene cluster. The glycosyl synthase, transfer, and regulatory genes of the upstream region of the meg PKS were contained in the nucleotide sequence which had a 9024 nucleotide sequence given in the specification. The glycosyl synthase, transfer, and regulatory genes of the downstream region of the meg PKS were contained in the nucleotide sequence which had a 17596 nucleotide sequence given in the specification. (51 pages)

ACCESSION NUMBER: 2004-10434 BIOTECHDS

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide; involving vector-mediated gene transfer and expression in host cell for polyketide production

AUTHOR: HUTCHINSON R C; KATZ L; REID R; HU Z; GRAMAO H

PATENT ASSIGNEE: KOSAN BIOSCIENCES INC

PATENT INFO: WO 2004003169 8 Jan 2004

APPLICATION INFO: WO 2003-US20681 30 Jun 2003

PRIORITY INFO: US 2002-393016 28 Jun 2002; US 2002-393016 28 Jun 2002

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: WPI: 2004-203379 [19]

L3 ANSWER 2 OF 27 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN 2004-203379 [19] WPIDS

AB WO2004003169 A UPAB: 20040318

NOVELTY - An isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

(1) an isolated, purified, or recombinant nucleic acid (II) comprising genes for the biosynthesis mycarose for attachment to a polyketide, the enzymes comprising the MegM, MegL, MegBIII, MegBIV, MegDIV, MegBII-2, and MegBVI enzymes;

(2) an isolated, purified, or recombinant nucleic acid (III) comprising genes for the biosynthesis mycarose for attachment of **negosamine** of a polyketide, the enzymes comprising the MegM, MegL, MegCII, MegBVI, MegDIV, MegDV, MegDII, MegDIII, and MegDI enzymes;

(3) an isolated, purified, or recombinant nucleic acid (IV) comprising genes for the biosynthesis of desosamine to a polyketide, the enzymes consisting of the MegM, MegL, MegCII, MegCIV, MegCV, MegDII, and MegDIII enzymes;

(4) an expression vector (V) comprising (I);

(5) a host cell comprising (I);

(6) a host cell comprising (II) that expresses a polyketide modifying enzyme encoded by a gene from a mycarose biosynthetic gene set, where the enzyme is chosen from MegM, MegL, MegBIII, MegBIV, MegDIV, MegBII-2, and MegBVI, MegBV, and MegF;

(7) a host cell comprising (III) that expresses a polyketide modifying enzyme encoded by a gene from a **megosamine** biosynthetic gene set, where the enzyme is chosen from MegM, MegL, MegCII, MegBVI, MegDIV, MegDV, MegDVI, MegDVI, MegDIII, MegDII, and MegDI; and

(8) a host cell comprising (IV) that expresses a polyketide modifying enzyme encoded by a gene from a desosamine biosynthetic gene set, where the enzyme is chosen from MegM, MegL, MegCII, MegCIV, MegCV, MegDII, and

MegDII, and MegCIII.

USE - (M1) is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising (I) under conditions in which the cell expresses a product of a gene encoded by (I) under conditions in which the unmodified polyketide is present, and producing the modified polyketide. In (M1), the cell further comprises (I) one or more module of a polyketide synthase. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, it its naturally occurring non-recombinant state cannot produce **megosamine**. (All claimed.)

DESCRIPTION OF DRAWING(S) - The drawing shows a schematic of the megalomicin polyketide synthase (meg DEBS) and corresponding meg genes upstream and downstream of the meg DEBS region and cosmids overlapping this region.

Dwg.1/3

ACCESSION NUMBER: 2004-203379 [19] WPIDS  
DOC. NO. CPI: C2004-080057  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
DERWENT CLASS: B03 B04 C02 D16  
INVENTOR(S): GRAMAJO, H; HU, Z; HUTCHINSON, R C; KATZ, L; REID, R; HUTCHINSON, C R  
PATENT ASSIGNEE(S): (KOSA-N) KOSAN BIOSCIENCES INC; (GRAM-I) GRAMAJO H; (HUZZ-I) HU Z; (HUTC-I) HUTCHINSON C R; (KATZ-I) KATZ L; (REID-I) REID R  
COUNTRY COUNT: 105  
PATENT INFORMATION:

PATENT NO	KIND	DATE	WEEK	LA	PG
WO 2004003169	A2	20040108 (200419)*	EN	51	
RW: AT BE BG CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE LS LU MC MW MZ NL OA PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW					
W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW					
AU 2003258978	A1	20040119 (200447)			
US 2004203015	A1	20041014 (200468)			

#### APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 2004003169	A2	WO 2003-US20681	20030630
AU 2003258978	A1	AU 2003-258978	20030630
US 2004203015	A1 Provisional	US 2002-393016P	20020628
		US 2003-611442	20030630

#### FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 2003258978	A1 Based on	WO 2004003169

PRIORITY APPLN. INFO: US 2002-393016P 20020628; US  
2003-611442 20030630

L3 ANSWER 3 OF 27 HCAPLUS COPYRIGHT 2006 ACS on STN  
TI Recombinant polynucleotides encoding megalomicin polyketide modifying

AB enzymes, and uses thereof  
The invention discloses biosynthetic, transfer and regulatory genes for various sugars to effectuate polyketide modification. It claims use of genes encoding biosynthetic enzymes for megalomicin A and use of genes involved in biosynthesis of thymine diphosphate-**megosamine**, TDP-mycarose, and TDP-desosamine. The genes that are claimed include megR, megF, **megK**, megCIV, megCV, megBVI, megBIII, megL, megM, megBIV, megDIV, megBII-2, megBV, megCII, megDV, megDII, megDIII, megDI, megCIII, megDVI, and megDVII. The invention further claims enzyme activities for attachment of mycarose, desosamine, and **megosamine** to a polyketide and the megF enzyme for hydroxylation of a polyketide. Materials and methods of the invention include heterologous promoters, expression vectors, recombinant host cells, and cell cultures to produce modified polyketides. Megalomicins are 6-O-glycosides of erythromycin C with acetyl or propionyl groups esterified to the 3''' or 4''' hydroxyls of the mycarose sugar. Their reported biol. activities include antibacterial activity, antiviral activity against herpes, and antiparasitic activity.

ACCESSION NUMBER: 2004:20831 HCAPLUS  
DOCUMENT NUMBER: 140:88769  
TITLE: Recombinant polynucleotides encoding megalomicin polyketide modifying enzymes, and uses thereof  
INVENTOR(S): Hutchinson, Richard C.; Katz, Leonard; Reid, Ralph; Hu, Zhihao; Gramajo, Hugo  
PATENT ASSIGNEE(S): Kosan Biosciences, Inc., USA  
SOURCE: PCT Int. Appl., 51 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004003169	A2	20040108	WO 2003-US20681	20030630
WO 2004003169	A3	20050804		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
AU 2003258978	A1	20040119	AU 2003-258978	20030630
US 2004203015	A1	20041014	US 2003-611442	20030630
PRIORITY APPLN. INFO.:			US 2002-393016P	P 20020628
			WO 2003-US20681	W 20030630

L3 ANSWER 4 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
AN ADI14170 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant

nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, it its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence is used in the exemplification of the invention.

ACCESSION NUMBER: ADI14170 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: Oligonucleotide of the invention SEQ ID NO:24.

L3 ANSWER 5 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14147 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, it its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence contains upstream megalomicin modification enzyme genes.

ACCESSION NUMBER: ADI14147 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea cosmid pKOS079-138B SEQ ID NO:1.

L3 ANSWER 6 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14161 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant

nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14161 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megDIV forward PCR primer SEQ ID NO:15.

L3 ANSWER 7 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14163 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14163 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBV forward PCR primer SEQ ID NO:17.

L3 ANSWER 8 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14157 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14157 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108

51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megBIV forward PCR primer SEQ ID NO:11.

L3 ANSWER 9 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14167 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14167 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108

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APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBVI forward PCR primer SEQ ID NO:21.

L3 ANSWER 10 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14155 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, it its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14155 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108

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APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megBIII forward PCR primer SEQ ID NO:9.

L3 ANSWER 11 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14151 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, it its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14151 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megL forward PCR primer SEQ ID NO:5.

L3 ANSWER 12 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14159 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14159 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megF forward PCR primer SEQ ID NO:13.

L3 ANSWER 13 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14160 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the

unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14160 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megF reverse PCR primer SEQ ID NO:14.

L3 ANSWER 14 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
AN ADI14166 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14166 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBII-2 reverse PCR primer SEQ ID NO:20.

L3 ANSWER 15 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
AN ADI14149 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant

nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence is used in the exemplification of the invention.

ACCESSION NUMBER: ADI14149 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: Synthetic oligonucleotide of the invention SEQ ID NO:3.

L3 ANSWER 16 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14154 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14154 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megM reverse PCR primer SEQ ID NO:8.

L3 ANSWER 17 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14158 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14158 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megBIV reverse PCR primer SEQ ID NO:12.

L3 ANSWER 18 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14156 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14156 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBIII reverse PCR primer SEQ ID NO:10.

L3 ANSWER 19 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14148 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, it its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence contains downstream megalomicin modification enzyme genes.

ACCESSION NUMBER: ADI14148 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea cosmid KOS205-57-2.3B SEQ ID NO:2.

L3 ANSWER 20 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14169 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, it its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence is used in the exemplification of the invention.

ACCESSION NUMBER: ADI14169 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: Oligonucleotide of the invention SEQ ID NO:23.

L3 ANSWER 21 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14168 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, it its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14168 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megBVI reverse PCR primer SEQ ID NO:22.

L3 ANSWER 22 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14165 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, it its naturally occurring

non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14165 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBII-2 forward PCR primer SEQ ID NO:19.

L3 ANSWER 23 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14153 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, it its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14153 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megM forward PCR primer SEQ ID NO:7.

L3 ANSWER 24 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14162 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A

method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14162 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megDIV reverse PCR primer SEQ ID NO:16.

L3 ANSWER 25 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14152 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14152 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megL reverse PCR primer SEQ ID NO:6.

L3 ANSWER 26 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme

e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14164 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14164 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megBV reverse PCR primer SEQ ID NO:18.

L3 ANSWER 27 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14150 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, **MegK**, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces **megosamine** and can attach **megosamine** to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce **megosamine**. The present sequence is used in the exemplification of the invention.

ACCESSION NUMBER: ADI14150 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: Synthetic oligonucleotide of the invention SEQ ID NO:4.

=> d his

(FILE 'HOME' ENTERED AT 12:24:36 ON 14 APR 2006)

FILE 'MEDLINE, BIOSIS, BIOTECHDS, EMBASE, WPIDS, HCAPLUS, DGENE,  
SCISEARCH' ENTERED AT 12:25:13 ON 14 APR 2006

L1 0 S MEGOSAMINE BIOSYNTHETIC GENE CLUSTER  
L2 52 S MEGOSAMINE  
L3 27 S L2 AND (MEGK)

=> e katz, l/au

E1 3 KATZ ZEILIG M/AU  
E2 18 KATZ ZVI/AU  
E3 0 --> KATZ, L/AU  
E4 1 KATZA E/AU  
E5 1 KATZAGIANNAKIS J/AU  
E6 1 KATZAKIAN/AU  
E7 9 KATZAKIAN A/AU  
E8 1 KATZAKIAN A J/AU  
E9 3 KATZAKIAN ARTHUR/AU  
E10 23 KATZAKIAN ARTHUR JR/AU  
E11 1 KATZAKIAN JUN ARTHUR/AU  
E12 1 KATZAKIAN JUNIOR ARTHUR/AU

=> e hutchinson, x/au

E1 1 HUTCHINSON YVONNE/AU  
E2 1 HUTCHINSON Z/AU  
E3 0 --> HUTCHINSON, X/AU  
E4 1 HUTCHINSONA W D/AU  
E5 1 HUTCHINSONCOLAS J/AU  
E6 1 HUTCHINSONCOLE H/AU  
E7 1 HUTCHINSONHOWORTH C/AU  
E8 1 HUTCHINSONI CLYDE A II/AU  
E9 2 HUTCHINSONWILLIAMS K/AU  
E10 7 HUTCHINSONWILLIAMS K A/AU  
E11 1 HUTCHINSRD/AU  
E12 1 HUTCHINSSO J T/AU

=> s megalomicin

L4 223 MEGALOMICIN

=> s 14 and gene

L5 104 L4 AND GENE

=> s 15 and (polyketide modifying enzyme)

L6 26 L5 AND (POLYKETIDE MODIFYING ENZYME)

=> d 16 ti abs ibib tot

L6 ANSWER 1 OF 26 BIOTECHDS COPYRIGHT 2006 THE THOMSON CORP. on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
polyketide modifying gene, there gene encodes  
polyketide modifying enzyme e.g., MegR, MegK,  
or MegM enzymes useful for producing modified polyketide;  
involving vector-mediated gene transfer and expression in  
host cell for polyketide production

AN 2004-10434 BIOTECHDS

AB DERVENT ABSTRACT:

NOVELTY - An isolated, purified, or recombinant nucleic acid (I)

comprising a polyketide modifying **gene**, where the **gene** encodes a **polyketide modifying enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following: (1) an isolated, purified, or recombinant nucleic acid (II) comprising genes for the biosynthesis mycarose for attachment to a polyketide, the enzymes comprising the MegM, MegL, MegBIII, MegBIV, MegDIV, MegBII-2, and MegBVI enzymes; (2) an isolated, purified, or recombinant nucleic acid (III) comprising genes for the biosynthesis mycarose for attachment of megosamine of a polyketide, the enzymes comprising the MegM, MegL, MegCII, MegBVI, MegDIV, MegDV, MegDII, MegDIII, and MegDI enzymes; (3) an isolated, purified, or recombinant nucleic acid (IV) comprising genes for the biosynthesis of desosamine to a polyketide, the enzymes consisting of the MegM, MegL, MegCII, MegCIV, MegCV, MegDII, and MegDIII enzymes; (4) an expression vector (V) comprising (I); (5) a host cell comprising (I); (6) a host cell comprising (II) that expresses a **polyketide modifying enzyme** encoded by a **gene** from a mycarose biosynthetic **gene** set, where the enzyme is chosen from MegM, MegL, MegBIII, MegBIV, MegDIV, MegBII-2, and MegBVI, MegBV, and MegF; (7) a host cell comprising (III) that expresses a **polyketide modifying enzyme** encoded by a **gene** from a megalomycin biosynthetic **gene** set, where the enzyme is chosen from MegM, MegL, MegCII, MegCIV, MegCV, MegDII, and MegDIII; and (8) a host cell comprising (IV) that expresses a **polyketide modifying enzyme** encoded by a **gene** from a desosamine biosynthetic **gene** set, where the enzyme is chosen from MegM, MegL, MegCII, MegCIV, MegCV, MegDII, and MegDIII.

BIOTECHNOLOGY - Preferred Nucleic Acid: In (I), the **gene** encodes a polyketide modifying enzymes chosen from MegR, MegK, MegCV, MegCIV, MegBVI, MegF, MegBII, MegM, and MegL. (I) further comprises **gene** encoding an enzyme for the attachment of mycarose to the polyketide, preferably MegBV enzyme. (I) further comprises **gene** encoding an enzyme for hydroxylation of the polyketide, preferably MegF enzyme. (IV) further comprises **gene** encoding an enzyme for the attachment of desosamine to the polyketide, preferably MegCIII enzyme. The polyketide modifying **gene** is operably linked to heterologous promoter.

USE - (M1) is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising (I) under conditions in which the cell expresses a product of a **gene** encoded by (I) under conditions in which the unmodified polyketide is present, and producing the modified polyketide. In (M1), the cell further comprises (I) one or more module of a polyketide synthase. The cell produces megalomycin and can attach megalomycin to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megalomycin. (All claimed.)

EXAMPLE - Isolation of the **megalomicin** biosynthetic **gene** cluster was as follows. A cosmid library was prepared in SuperCos vectors from Micromonospora megalomicea total DNA partially digested with Sau3AI and introduced into Escherichia coli using a Gigapack III XL in vitro packaging kit. 32P-labeled DNA probes encompassing the KS2 domain from DEBS, or a mixture of segments encompassing modules 1 and 2 from DEBS, were used separately to screen the cosmid library by colony hybridization. Several colonies which hybridized with the probes were further analyzed by sequencing the ends of their cosmid inserts using T3 and T7 primers. BLAST analysis of the sequences revealed several colonies with DNA sequences highly homologous to genes from the ery cluster. Together with restriction analysis, that led to the isolation of two overlapping cosmids, pKOS079-93A and pKOS079-93D which covered 45 kbase the meg cluster. A 400 base pair PCR fragment was generated from the left end of pKOS079-93D and used to

reprobe the cosmid library. Likewise, a 200 base pair PCR fragment generated from the right end of pKOSO79-93A was used to reprobe the cosmid library. Analysis of hybridizing colonies, as described above, resulted in identification of two additional cosmids pKOSO79-93B which adjacent to the 5' end of pKOSO79-93D and pKOSO5.57-2.3B which overlapped the 3' ends of pKOSO79-93A and pKOSO79-93D cosmids. BLAST analysis of the far left and right end sequences of these cosmids indicated no homology to any known genes related to polyketide biosynthesis, and therefore indicates that the set of four cosmids spans the entire **megalomicin** biosynthetic **gene** cluster. The glycosyl synthase, transfer, and regulatory genes of the upstream region of the meg PKS were contained in the nucleotide sequence which had a 9024 nucleotide sequence given in the specification. The glycosyl synthase, transfer, and regulatory genes of the downstream region of the meg PKS were contained in the nucleotide sequence which had a 17596 nucleotide sequence given in the specification. (51 pages)

ACCESSION NUMBER: 2004-10434 BIOTECHDS

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, where **gene** encodes **Polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide; involving vector-mediated **gene** transfer and expression in host cell for polyketide production

AUTHOR: HUTCHINSON R C; KATZ L; REID R; HU Z; GRAMAO H

PATENT ASSIGNEE: KOSAN BIOSCIENCES INC

PATENT INFO: WO 2004003169 8 Jan 2004

APPLICATION INFO: WO 2003-US20681 30 Jun 2003

PRIORITY INFO: US 2002-393016 28 Jun 2002; US 2002-393016 28 Jun 2002

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: WPI: 2004-203379 [19]

L6 ANSWER 2 OF 26 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, where **gene** encodes **Polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

AN 2004-203379 [19] WPIDS

AB WO2004003169 A UPAB: 20040318

NOVELTY - An isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying **gene**, where the **gene** encodes a **Polyketide modifying enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

(1) an isolated, purified, or recombinant nucleic acid (II) comprising genes for the biosynthesis mycarose for attachment to a polyketide, the enzymes comprising the MegM, MegL, MegBIII, MegBIV, MegDIV, MegBII-2, and MegBVI enzymes;

(2) an isolated, purified, or recombinant nucleic acid (III) comprising genes for the biosynthesis mycarose for attachment of megalamic acid to a polyketide, the enzymes comprising the MegM, MegL, MegCII, MegBVI, MegDIV, MegDV, MegDII, MegDIII, and MegDI enzymes;

(3) an isolated, purified, or recombinant nucleic acid (IV) comprising genes for the biosynthesis of desosamine to a polyketide, the enzymes consisting of the MegM, MegL, MegCII, MegCIV, MegCV, MegDII, and MegDIII enzymes;

(4) an expression vector (V) comprising (I);

(5) a host cell comprising (I);

(6) a host cell comprising (II) that expresses a **Polyketide modifying enzyme** encoded by a **gene** from a

mycarose biosynthetic **gene** set, where the enzyme is chosen from MegM, MegL, MegBIII, MegBIV, MegDIV, MegBII-2, and MegBVI, MegBV, and MegF;

(7) a host cell comprising (III) that expresses a **polyketide modifying enzyme** encoded by a **gene** from a megalosamine biosynthetic **gene** set, where the enzyme is chosen from MegM, MegL, MegCII, MegBVI, MegDIV, MegDV, MegDVI, MegDVII, MegDII, MegDIII, and MegDI; and

(8) a host cell comprising (IV) that expresses a **polyketide modifying enzyme** encoded by a **gene** from a desosamine biosynthetic **gene** set, where the enzyme is chosen from MegM, MegL, MegCII, MegCIV, MegCV, MegDII, and MegCIII.

USE - (M1) is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising (I) under conditions in which the cell expresses a product of a **gene** encoded by (I) under conditions in which the unmodified polyketide is present, and producing the modified polyketide. In (M1), the cell further comprises (I) one or more module of a polyketide synthase. The cell produces megalosamine and can attach megalosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megalosamine. (All claimed.)

DESCRIPTION OF DRAWING(S) - The drawing shows a schematic of the **megalomicin** polyketide synthase (meg DEBS) and corresponding meg genes upstream and downstream of the meg DEBS region and cosmids overlapping this region.

Dwg.1/3

ACCESSION NUMBER: 2004-203379 [19] WPIDS  
DOC. NO. CPI: C2004-080057  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, where **gene** encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
DERWENT CLASS: B03 B04 C02 D16  
INVENTOR(S): GRAMAJO, H; HU, Z; HUTCHINSON, R C; KATZ, L; REID, R; HUTCHINSON, C R  
PATENT ASSIGNEE(S): (KOSA-N) KOSAN BIOSCIENCES INC; (GRAM-I) GRAMAJO H; (HUZZ-I) HU Z; (HUTC-I) HUTCHINSON C R; (KATZ-I) KATZ L; (REID-I) REID R  
COUNTRY COUNT: 105  
PATENT INFORMATION:

PATENT NO	KIND	DATE	WEEK	LA	PG
<hr/>					
WO 2004003169	A2	20040108 (200419)*	EN	51	
RW:	AT BE BG CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE LS LU MC MW MZ NL OA PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW				
W:	AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW				
AU 2003258978	A1	20040119 (200447)			
US 2004203015	A1	20041014 (200468)			

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
<hr/>			
WO 2004003169	A2	WO 2003-US20681	20030630
AU 2003258978	A1	AU 2003-258978	20030630
US 2004203015	A1 Provisional	US 2002-393016P	20020628
		US 2003-611442	20030630

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 2003258978	A1 Based on	WO 2004003169

PRIORITY APPLN. INFO: US 2002-393016P 20020628; US  
2003-611442 20030630

L6 ANSWER 3 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
 TI Novel isolated, purified, or recombinant nucleic acid comprising  
 polyketide modifying gene, there **gene** encodes  
**polyketide modifying enzyme** e.g., MegR, MegK,  
 or MegM enzymes useful for producing modified polyketide.  
 AN ADI14170 DNA DGENE  
 AB The invention relates to a novel isolated, purified, or recombinant  
 nucleic acid (I) comprising a polyketide modifying **gene**, where  
 the **gene** encodes a **polyketide modifying enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI,  
 MegBIII, MegL, and MegM enzymes. A method of the invention is useful for  
 producing a modified polyketide, which involves culturing a recombinant  
 cell comprising the recombinant nucleic acid under conditions in which  
 the cell expresses a product of a **gene** encoded by the nucleic  
 acid under conditions in which the unmodified polyketide is present, and  
 producing the modified polyketide. The cell produces megosamine and can  
 attach megosamine to a polyketide, where the cell, it its naturally  
 occurring non-recombinant state cannot produce megosamine. The present  
 sequence is used in the exemplification of the invention.

ACCESSION NUMBER: ADI14170 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid  
 comprising polyketide modifying **gene**, there  
**gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for  
 producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: Oligonucleotide of the invention SEQ ID NO:24.

L6 ANSWER 4 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
 TI Novel isolated, purified, or recombinant nucleic acid comprising  
 polyketide modifying gene, there **gene** encodes  
**polyketide modifying enzyme** e.g., MegR, MegK,  
 or MegM enzymes useful for producing modified polyketide.

AN ADI14147 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant  
 nucleic acid (I) comprising a polyketide modifying **gene**, where  
 the **gene** encodes a **polyketide modifying enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI,  
 MegBIII, MegL, and MegM enzymes. A method of the invention is useful for  
 producing a modified polyketide, which involves culturing a recombinant  
 cell comprising the recombinant nucleic acid under conditions in which  
 the cell expresses a product of a **gene** encoded by the nucleic  
 acid under conditions in which the unmodified polyketide is present, and  
 producing the modified polyketide. The cell produces megosamine and can  
 attach megosamine to a polyketide, where the cell, it its naturally  
 occurring non-recombinant state cannot produce megosamine. The present

sequence contains upstream **megalomicin** modification enzyme genes.

ACCESSION NUMBER: ADI14147 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, there **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea cosmid pKOS079-138B SEQ ID NO:1.

L6 ANSWER 5 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, there **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
AN ADI14161 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying **gene**, where the **gene** encodes a **polyketide modifying enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a **gene** encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, it its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a **gene** of the invention.

ACCESSION NUMBER: ADI14161 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, there **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megDIV forward PCR primer SEQ ID NO:15.

L6 ANSWER 6 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, there **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
AN ADI14163 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying **gene**, where the **gene** encodes a **polyketide modifying**

**enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a **gene** encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a **gene** of the invention.

ACCESSION NUMBER: ADI14163 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, where **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBV forward PCR primer SEQ ID NO:17.

L6 ANSWER 7 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, where **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
AN ADI14157 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying **gene**, where the **gene** encodes a **polyketide modifying enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a **gene** encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a **gene** of the invention.

ACCESSION NUMBER: ADI14157 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, where **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBIV forward PCR primer SEQ ID NO:11.

L6 ANSWER 8 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
polyketide modifying gene, there gene encodes  
**polyketide modifying enzyme** e.g., MegR, MegK,  
or MegM enzymes useful for producing modified polyketide.  
AN ADI14167 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant  
nucleic acid (I) comprising a polyketide modifying gene, where  
the gene encodes a **polyketide modifying**  
**enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI,  
MegBIII, MegL, and MegM enzymes. A method of the invention is useful for  
producing a modified polyketide, which involves culturing a recombinant  
cell comprising the recombinant nucleic acid under conditions in which  
the cell expresses a product of a gene encoded by the nucleic  
acid under conditions in which the unmodified polyketide is present, and  
producing the modified polyketide. The cell produces megosamine and can  
attach megosamine to a polyketide, where the cell, in its naturally  
occurring non-recombinant state cannot produce megosamine. The present  
sequence represents a PCR primer used to amplify a gene of the  
invention.

ACCESSION NUMBER: ADI14167 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid  
comprising polyketide modifying gene, there  
gene encodes **polyketide modifying**  
**enzyme** e.g., MegR, MegK, or MegM enzymes useful for  
producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBVI forward PCR primer SEQ ID NO:21.

L6 ANSWER 9 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
polyketide modifying gene, there gene encodes  
**polyketide modifying enzyme** e.g., MegR, MegK,  
or MegM enzymes useful for producing modified polyketide.  
AN ADI14155 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant  
nucleic acid (I) comprising a polyketide modifying gene, where  
the gene encodes a **polyketide modifying**  
**enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI,  
MegBIII, MegL, and MegM enzymes. A method of the invention is useful for  
producing a modified polyketide, which involves culturing a recombinant  
cell comprising the recombinant nucleic acid under conditions in which  
the cell expresses a product of a gene encoded by the nucleic  
acid under conditions in which the unmodified polyketide is present, and  
producing the modified polyketide. The cell produces megosamine and can  
attach megosamine to a polyketide, where the cell, in its naturally  
occurring non-recombinant state cannot produce megosamine. The present  
sequence represents a PCR primer used to amplify a gene of the  
invention.

ACCESSION NUMBER: ADI14155 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid  
comprising polyketide modifying gene, there  
gene encodes **polyketide modifying**  
**enzyme** e.g., MegR, MegK, or MegM enzymes useful for  
producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBIII forward PCR primer SEQ ID NO:9.

L6 ANSWER 10 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
polyketide modifying gene, there **gene** encodes  
**polyketide modifying enzyme** e.g., MegR, MegK,  
or MegM enzymes useful for producing modified polyketide.  
AN ADI14151 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant  
nucleic acid (I) comprising a polyketide modifying **gene**, where  
the **gene** encodes a **polyketide modifying**  
**enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI,  
MegBIII, MegL, and MegM enzymes. A method of the invention is useful for  
producing a modified polyketide, which involves culturing a recombinant  
cell comprising the recombinant nucleic acid under conditions in which  
the cell expresses a product of a **gene** encoded by the nucleic  
acid under conditions in which the unmodified polyketide is present, and  
producing the modified polyketide. The cell produces megosamine and can  
attach megosamine to a polyketide, where the cell, it its naturally  
occurring non-recombinant state cannot produce megosamine. The present  
sequence represents a PCR primer used to amplify a **gene** of the  
invention.

ACCESSION NUMBER: ADI14151 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid  
comprising polyketide modifying **gene**, there  
**gene** encodes **polyketide modifying**  
**enzyme** e.g., MegR, MegK, or MegM enzymes useful for  
producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megL forward PCR primer SEQ ID NO:5.

L6 ANSWER 11 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
polyketide modifying gene, there **gene** encodes  
**polyketide modifying enzyme** e.g., MegR, MegK,  
or MegM enzymes useful for producing modified polyketide.  
AN ADI14159 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant  
nucleic acid (I) comprising a polyketide modifying **gene**, where  
the **gene** encodes a **polyketide modifying**  
**enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI,  
MegBIII, MegL, and MegM enzymes. A method of the invention is useful for  
producing a modified polyketide, which involves culturing a recombinant  
cell comprising the recombinant nucleic acid under conditions in which  
the cell expresses a product of a **gene** encoded by the nucleic  
acid under conditions in which the unmodified polyketide is present, and  
producing the modified polyketide. The cell produces megosamine and can  
attach megosamine to a polyketide, where the cell, it its naturally  
occurring non-recombinant state cannot produce megosamine. The present

sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14159 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megF forward PCR primer SEQ ID NO:13.

L6 ANSWER 12 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

AN ADI14160 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, it its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14160 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megF reverse PCR primer SEQ ID NO:14.

L6 ANSWER 13 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

AN ADI14166 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying

**enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a **gene** encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a **gene** of the invention.

ACCESSION NUMBER: ADI14166 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, where **gene** encodes polyketide modifying **enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBII-2 reverse PCR primer SEQ ID NO:20.

L6 ANSWER 14 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, where **gene** encodes polyketide modifying **enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
AN ADI14149 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying **gene**, where the **gene** encodes a polyketide modifying **enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a **gene** encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence is used in the exemplification of the invention.

ACCESSION NUMBER: ADI14149 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, where **gene** encodes polyketide modifying **enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: Synthetic oligonucleotide of the invention SEQ ID NO:3.

L6 ANSWER 15 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

AN ADI14154 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megalomicea and can attach megalomicea to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megalomicea. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14154 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megM reverse PCR primer SEQ ID NO:8.

L6 ANSWER 16 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

AN ADI14158 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megalomicea and can attach megalomicea to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megalomicea. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14158 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, there gene encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBIV reverse PCR primer SEQ ID NO:12.

L6 ANSWER 17 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
polyketide modifying gene, there **gene** encodes  
**polyketide modifying enzyme** e.g., MegR, MegK,  
or MegM enzymes useful for producing modified polyketide.  
AN ADI14156 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant  
nucleic acid (I) comprising a polyketide modifying **gene**, where  
the **gene** encodes a **polyketide modifying**  
**enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI,  
MegBIII, MegL, and MegM enzymes. A method of the invention is useful for  
producing a modified polyketide, which involves culturing a recombinant  
cell comprising the recombinant nucleic acid under conditions in which  
the cell expresses a product of a **gene** encoded by the nucleic  
acid under conditions in which the unmodified polyketide is present, and  
producing the modified polyketide. The cell produces megosamine and can  
attach megosamine to a polyketide, where the cell, it its naturally  
occurring non-recombinant state cannot produce megosamine. The present  
sequence represents a PCR primer used to amplify a **gene** of the  
invention.

ACCESSION NUMBER: ADI14156 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid  
comprising polyketide modifying **gene**, there  
**gene** encodes **polyketide modifying**  
**enzyme** e.g., MegR, MegK, or MegM enzymes useful for  
producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBIII reverse PCR primer SEQ ID NO:10.

L6 ANSWER 18 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
polyketide modifying gene, there **gene** encodes  
**polyketide modifying enzyme** e.g., MegR, MegK,  
or MegM enzymes useful for producing modified polyketide.  
AN ADI14148 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant  
nucleic acid (I) comprising a polyketide modifying **gene**, where  
the **gene** encodes a **polyketide modifying**  
**enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI,  
MegBIII, MegL, and MegM enzymes. A method of the invention is useful for  
producing a modified polyketide, which involves culturing a recombinant  
cell comprising the recombinant nucleic acid under conditions in which  
the cell expresses a product of a **gene** encoded by the nucleic  
acid under conditions in which the unmodified polyketide is present, and  
producing the modified polyketide. The cell produces megosamine and can  
attach megosamine to a polyketide, where the cell, it its naturally  
occurring non-recombinant state cannot produce megosamine. The present  
sequence contains downstream **megalomicin** modification enzyme

genes.

ACCESSION NUMBER: ADI14148 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, there **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea cosmid KOS205-57-2.3B SEQ ID NO:2.

L6 ANSWER 19 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, there **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

AN ADI14169 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying **gene**, where the **gene** encodes a **polyketide modifying enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a **gene** encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence is used in the exemplification of the invention.

ACCESSION NUMBER: ADI14169 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, there **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: Oligonucleotide of the invention SEQ ID NO:23.

L6 ANSWER 20 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, there **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

AN ADI14168 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying **gene**, where the **gene** encodes a **polyketide modifying enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for

producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a **gene** encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a **gene** of the invention.

ACCESSION NUMBER: ADI14168 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, where **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBVI reverse PCR primer SEQ ID NO:22.

L6 ANSWER 21 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, where **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
AN ADI14165 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying **gene**, where the **gene** encodes a **polyketide modifying enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a **gene** encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a **gene** of the invention.

ACCESSION NUMBER: ADI14165 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, where **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBII-2 forward PCR primer SEQ ID NO:19.

L6 ANSWER 22 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising

polyketide modifying gene, there **gene** encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

AN ADI14153 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the **gene** encodes a polyketide modifying enzyme chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a **gene** encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a **gene** of the invention.

ACCESSION NUMBER: ADI14153 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where **gene** encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108

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APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megM forward PCR primer SEQ ID NO:7.

L6 ANSWER 23 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where **gene** encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

AN ADI14162 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the **gene** encodes a polyketide modifying enzyme chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a **gene** encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a **gene** of the invention.

ACCESSION NUMBER: ADI14162 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where **gene** encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108

51

APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megDIV reverse PCR primer SEQ ID NO:16.

L6 ANSWER 24 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
AN ADI14152 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying **gene**, where the **gene** encodes a **polyketide modifying enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a **gene** encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a **gene** of the invention.

ACCESSION NUMBER: ADI14152 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying **gene**, where **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megL reverse PCR primer SEQ ID NO:6.

L6 ANSWER 25 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where **gene** encodes **polyketide modifying enzyme** e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
AN ADI14164 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying **gene**, where the **gene** encodes a **polyketide modifying enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a **gene** encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a **gene** of the invention.

ACCESSION NUMBER: ADI14164 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBV reverse PCR primer SEQ ID NO:18.

L6 ANSWER 26 OF 26 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
AN ADI14150 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a polyketide modifying gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence is used in the exemplification of the invention.

ACCESSION NUMBER: ADI14150 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising polyketide modifying gene, where gene encodes polyketide modifying enzyme e.g., MegR, MegK, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: Synthetic oligonucleotide of the invention SEQ ID NO:4.

# Refine Search

## Search Results -

Terms	Documents
L3 and (megF or megK or megcIV or megBVI)	1

Database:

US Pre-Grant Publication Full-Text Database  
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L4

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DATE: Friday, April 14, 2006 [Printable Copy](#) [Create Case](#)

Set Name Query  
side by side

Hit Count Set Name  
result set

DB=USPT; PLUR=YES; OP=OR

<u>L4</u>	L3 and (megF or megK or megcIV or megBVI)	1	<u>L4</u>
<u>L3</u>	6524841.pn.	1	<u>L3</u>
<u>L2</u>	6303342.pn.	1	<u>L2</u>
<u>L1</u>	5998194.pn.	1	<u>L1</u>

END OF SEARCH HISTORY

# Refine Search

## Search Results -

Terms	Documents
L9 and L8	0

Database:

US Pre-Grant Publication Full-Text Database  
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IBM Technical Disclosure Bulletins

Search:

L10

Refine Search

Recall Text

Clear

Interrupt

## Search History

DATE: Friday, April 14, 2006 [Printable Copy](#) [Create Case](#)

### Set Name Query

side by side

DB=USPT; PLUR=YES; OP=OR

Hit Count Set Name  
result set

<u>L10</u>	L9 and l8	0	<u>L10</u>
<u>L9</u>	hutchinson.in.	1326	<u>L9</u>
<u>L8</u>	katz.in.	2281	<u>L8</u>
<u>L7</u>	L1 and (method of producing modified polyketide)	1	<u>L7</u>
<u>L6</u>	L4 and (Host cell)	1	<u>L6</u>
<u>L5</u>	(L4 and Host cell)	506752	<u>L5</u>
<u>L4</u>	L3 and (megF or megK or megcIV or megBVI)	1	<u>L4</u>
<u>L3</u>	6524841.pn.	1	<u>L3</u>
<u>L2</u>	6303342.pn.	1	<u>L2</u>
<u>L1</u>	5998194.pn.	1	<u>L1</u>

END OF SEARCH HISTORY

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INPADOC  
NEWS 6 JAN 17 Pre-1988 INPI data added to MARPAT  
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property data  
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NEWS 17 MAR 03 Updates in PATDPA; addition of IPC 8 data without attributes  
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thesaurus added in PCTFULL  
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in MARPAT  
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second quarter; strategies may be affected  
  
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=> s polyketide modifying gene  
L1 27 POLYKETIDE MODIFYING GENE

=> s 11 and enzyme  
L2 27 L1 AND ENZYME

=> s 12 and (MegF or MegK or MegCIV or MegCV or MegBIII)  
L3 27 L2 AND (MEGF OR ME GK OR MEGCIV OR MEGCV OR MEGBIII)

=> d 13 ti abs ibib tot

L3 ANSWER 1 OF 27 USPATFULL on STN

## TI Recombinant genes for polyketide modifying enzymes

**AB** Materials and methods to produce modified polyketides are disclosed. The biosynthesis, transfer and regulator genes for various sugars to effectuate polyketide modification are disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

THIS INDEXING IS AVAILABLE FOR THIS PATENT.

TITLE: Recombinant genes for polyketide modifying enzymes  
INVENTOR(S): Hutchinson, C. Richard, San Mateo, CA, UNITED STATES  
Katz, Leonard, Oakland, CA, UNITED STATES  
Reid, Ralph, San Rafael, CA, UNITED STATES  
Hu, Zhihao, Castro Valley, CA, UNITED STATES  
Gramajo, Hugo, Berkeley, CA, UNITED STATES

	NUMBER	KIND	DATE
PATENT INFORMATION:	US 2004203015	A1	20041014
APPLICATION INFO.:	US 2003-611442	A1	20030630 (10)

	NUMBER	DATE
PRIORITY INFORMATION:	US 2002-393016P	20020628 (60)
DOCUMENT TYPE:	Utility	
FILE SEGMENT:	APPLICATION	
LEGAL REPRESENTATIVE:	Ted Apple, (Townsend and Townsend and Crew), 379 Lytton Avenue, Palo Alto, CA, 94301	
NUMBER OF CLAIMS:	23	
EXEMPLARY CLAIM:	1	
NUMBER OF DRAWINGS:	3 Drawing Page(s)	
LINE COUNT:	2721	

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 2 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
**polyketide modifying gene**, where gene encodes  
polyketide modifying enzyme e.g., MegR, **MegK**, or MegM  
enzymes useful for producing modified polyketide.  
AN ADI14170 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying enzyme chosen from MegR, **MegF**, **MegK**,  
**MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, **MegL**, and  
MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megalosamine and can attach megalosamine to a polyketide, where the cell, if its naturally occurring non-recombinant state cannot produce megalosamine. The present sequence is used in the exemplification of the invention.

ACCESSION NUMBER: ADI14170 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: Oligonucleotide of the invention SEQ ID NO:24.

L3 ANSWER 3 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising

**polyketide modifying gene**, there gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14147 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying enzyme chosen from MegR, **MegF**, **MegK**, **MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence contains upstream megalomicin modification **enzyme** genes.

ACCESSION NUMBER: ADI14147 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, there gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea cosmid pKOS079-138B SEQ ID NO:1.

L3 ANSWER 4 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, there gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
AN ADI14161 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying enzyme chosen from MegR, **MegF**, **MegK**, **MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14161 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, there gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megDIV forward PCR primer SEQ ID NO:15.

L3 ANSWER 5 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
**polyketide modifying gene**, there gene encodes  
polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM  
enzymes useful for producing modified polyketide.  
AN ADI14163 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant  
nucleic acid (I) comprising a **polyketide modifying**  
**gene**, where the gene encodes a polyketide modifying  
**enzyme** chosen from MegR, **MegF**, **MegK**,  
**MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, **MegL**, and  
MegM enzymes. A method of the invention is useful for producing a  
modified polyketide, which involves culturing a recombinant cell  
comprising the recombinant nucleic acid under conditions in which the  
cell expresses a product of a gene encoded by the nucleic acid under  
conditions in which the unmodified polyketide is present, and producing  
the modified polyketide. The cell produces megosamine and can attach  
megosamine to a polyketide, where the cell, it its naturally occurring  
non-recombinant state cannot produce megosamine. The present sequence  
represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14163 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid  
comprising **polyketide modifying**  
**gene**, there gene encodes polyketide modifying  
**enzyme** e.g., MegR, **MegK**, or MegM enzymes  
useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBV forward PCR primer SEQ ID NO:17.

L3 ANSWER 6 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
**polyketide modifying gene**, there gene encodes  
polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM  
enzymes useful for producing modified polyketide.  
AN ADI14157 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant  
nucleic acid (I) comprising a **polyketide modifying**  
**gene**, where the gene encodes a polyketide modifying  
**enzyme** chosen from MegR, **MegF**, **MegK**,  
**MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, **MegL**, and  
MegM enzymes. A method of the invention is useful for producing a  
modified polyketide, which involves culturing a recombinant cell  
comprising the recombinant nucleic acid under conditions in which the  
cell expresses a product of a gene encoded by the nucleic acid under  
conditions in which the unmodified polyketide is present, and producing  
the modified polyketide. The cell produces megosamine and can attach  
megosamine to a polyketide, where the cell, it its naturally occurring  
non-recombinant state cannot produce megosamine. The present sequence  
represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14157 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBIV forward PCR primer SEQ ID NO:11.

L3 ANSWER 7 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
AN ADI14167 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying **enzyme** chosen from MegR, **MegF**, **MegK**, **MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14167 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBVI forward PCR primer SEQ ID NO:21.

L3 ANSWER 8 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
AN ADI14155 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying **enzyme** chosen from MegR, **MegF**, **MegK**, **MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, MegL, and

MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megalomicea and can attach megalomicea to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megalomicea. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14155 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea **megBIII** forward PCR primer SEQ ID NO:9.

L3 ANSWER 9 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
AN ADI14151 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying enzyme chosen from MegR, **MegF**, **MegK**, **MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megalomicea and can attach megalomicea to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megalomicea. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14151 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megL forward PCR primer SEQ ID NO:5.

L3 ANSWER 10 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14159 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying enzyme chosen from MegR, **MegF**, **MegK**, **MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megalomicea and can attach megalomicea to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megalomicea. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14159 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea **megF** forward PCR primer SEQ ID NO:13..

L3 ANSWER 11 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN

TI Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14160 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying enzyme chosen from MegR, **MegF**, **MegK**, **MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, MegL, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megalomicea and can attach megalomicea to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megalomicea. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14160 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megF reverse PCR primer SEQ ID NO:14.

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L3 ANSWER 12 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
**polyketide modifying gene**, where gene encodes  
polyketide modifying enzyme e.g., MegR, **MegK**, or MegM  
enzymes useful for producing modified polyketide.  
AN ADI14166 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant  
nucleic acid (I) comprising a **polyketide modifying**  
**gene**, where the gene encodes a polyketide modifying  
enzyme chosen from MegR, **MegF**, **MegK**,  
**MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, MegL, and  
MegM enzymes. A method of the invention is useful for producing a  
modified polyketide, which involves culturing a recombinant cell  
comprising the recombinant nucleic acid under conditions in which the  
cell expresses a product of a gene encoded by the nucleic acid under  
conditions in which the unmodified polyketide is present, and producing  
the modified polyketide. The cell produces megosamine and can attach  
megosamine to a polyketide, where the cell, in its naturally occurring  
non-recombinant state cannot produce megosamine. The present sequence  
represents a PCR primer used to amplify a gene of the invention.  
ACCESSION NUMBER: ADI14166 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid  
comprising **polyketide modifying**  
**gene**, where gene encodes polyketide modifying  
enzyme e.g., MegR, **MegK**, or MegM enzymes  
useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBII-2 reverse PCR primer SEQ ID NO:20.

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L3 ANSWER 13 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
**polyketide modifying gene**, where gene encodes  
polyketide modifying enzyme e.g., MegR, **MegK**, or MegM  
enzymes useful for producing modified polyketide.  
AN ADI14149 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant  
nucleic acid (I) comprising a **polyketide modifying**  
**gene**, where the gene encodes a polyketide modifying  
enzyme chosen from MegR, **MegF**, **MegK**,  
**MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, MegL, and  
MegM enzymes. A method of the invention is useful for producing a  
modified polyketide, which involves culturing a recombinant cell  
comprising the recombinant nucleic acid under conditions in which the  
cell expresses a product of a gene encoded by the nucleic acid under  
conditions in which the unmodified polyketide is present, and producing  
the modified polyketide. The cell produces megosamine and can attach

megosamine to a polyketide, where the cell, it its naturally occurring non-recombinant state cannot produce megosamine. The present sequence is used in the exemplification of the invention.

ACCESSION NUMBER: ADI14149 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, there gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: Synthetic oligonucleotide of the invention SEQ ID NO:3.

L3 ANSWER 14 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, there gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
AN ADI14154 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying **enzyme** chosen from MegR, **MegF**, **MegK**, **MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, **MegL**, and **MegM** enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, it its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14154 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, there gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megM reverse PCR primer SEQ ID NO:8.

L3 ANSWER 15 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, there gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
AN ADI14158 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying**

gene, where the gene encodes a polyketide modifying enzyme chosen from MegR, **MegF**, **MegK**, **MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, **MegL**, and **MegM** enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14158 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megBIV reverse PCR primer SEQ ID NO:12.

L3 ANSWER 16 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14156 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying **enzyme** chosen from MegR, **MegF**, **MegK**, **MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, **MegL**, and **MegM** enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14156 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea **megBIII** reverse PCR primer SEQ ID

NO:10.

L3 ANSWER 17 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
**polyketide modifying gene**, where gene encodes  
polyketide modifying enzyme e.g., MegR, **MegK**, or MegM  
enzymes useful for producing modified polyketide.  
AN ADI14148 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant  
nucleic acid (I) comprising a **polyketide modifying**  
**gene**, where the gene encodes a polyketide modifying  
enzyme chosen from MegR, **MegF**, **MegK**,  
**MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, MegL, and  
MegM enzymes. A method of the invention is useful for producing a  
modified polyketide, which involves culturing a recombinant cell  
comprising the recombinant nucleic acid under conditions in which the  
cell expresses a product of a gene encoded by the nucleic acid under  
conditions in which the unmodified polyketide is present, and producing  
the modified polyketide. The cell produces megalomycin and can attach  
megalomycin to a polyketide, where the cell, in its naturally occurring  
non-recombinant state cannot produce megalomycin. The present sequence  
contains downstream megalomycin modification **enzyme** genes.  
ACCESSION NUMBER: ADI14148 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid  
comprising **polyketide modifying**  
**gene**, where gene encodes polyketide modifying  
enzyme e.g., MegR, **MegK**, or MegM enzymes  
useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea cosmid KOS205-57-2.3B SEQ ID NO:2.

L3 ANSWER 18 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
**polyketide modifying gene**, where gene encodes  
polyketide modifying enzyme e.g., MegR, **MegK**, or MegM  
enzymes useful for producing modified polyketide.  
AN ADI14169 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant  
nucleic acid (I) comprising a **polyketide modifying**  
**gene**, where the gene encodes a polyketide modifying  
enzyme chosen from MegR, **MegF**, **MegK**,  
**MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, MegL, and  
MegM enzymes. A method of the invention is useful for producing a  
modified polyketide, which involves culturing a recombinant cell  
comprising the recombinant nucleic acid under conditions in which the  
cell expresses a product of a gene encoded by the nucleic acid under  
conditions in which the unmodified polyketide is present, and producing  
the modified polyketide. The cell produces megalomycin and can attach  
megalomycin to a polyketide, where the cell, in its naturally occurring  
non-recombinant state cannot produce megalomycin. The present sequence is  
used in the exemplification of the invention.  
ACCESSION NUMBER: ADI14169 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid  
comprising **polyketide modifying**  
**gene**, where gene encodes polyketide modifying  
enzyme e.g., MegR, **MegK**, or MegM enzymes

useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: Oligonucleotide of the invention SEQ ID NO:23.

L3 ANSWER 19 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
**polyketide modifying gene**, where gene encodes  
polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM  
enzymes useful for producing modified polyketide.  
AN ADI14168 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant  
nucleic acid (I) comprising a **polyketide modifying**  
**gene**, where the gene encodes a polyketide modifying  
**enzyme** chosen from MegR, **MegF**, **MegK**,  
**MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, **MegL**, and  
MegM enzymes. A method of the invention is useful for producing a  
modified polyketide, which involves culturing a recombinant cell  
comprising the recombinant nucleic acid under conditions in which the  
cell expresses a product of a gene encoded by the nucleic acid under  
conditions in which the unmodified polyketide is present, and producing  
the modified polyketide. The cell produces megalomycin and can attach  
megalomycin to a polyketide, where the cell, in its naturally occurring  
non-recombinant state cannot produce megalomycin. The present sequence  
represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14168 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid  
comprising **polyketide modifying**  
**gene**, where gene encodes polyketide modifying  
**enzyme** e.g., MegR, **MegK**, or MegM enzymes  
useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megBVI reverse PCR primer SEQ ID NO:22.

L3 ANSWER 20 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
**polyketide modifying gene**, where gene encodes  
polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM  
enzymes useful for producing modified polyketide.  
AN ADI14165 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant  
nucleic acid (I) comprising a **polyketide modifying**  
**gene**, where the gene encodes a polyketide modifying  
**enzyme** chosen from MegR, **MegF**, **MegK**,  
**MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, **MegL**, and  
MegM enzymes. A method of the invention is useful for producing a  
modified polyketide, which involves culturing a recombinant cell  
comprising the recombinant nucleic acid under conditions in which the  
cell expresses a product of a gene encoded by the nucleic acid under  
conditions in which the unmodified polyketide is present, and producing

the modified polyketide. The cell produces megalomicea and can attach megalomicea to a polyketide, where the cell, it its naturally occurring non-recombinant state cannot produce megalomicea. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14165 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, there gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megBII-2 forward PCR primer SEQ ID NO:19.

L3 ANSWER 21 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, there gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14153 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying **enzyme** chosen from MegR, **MegF**, **MegK**, **MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, **MegL**, and MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megalomicea and can attach megalomicea to a polyketide, where the cell, it its naturally occurring non-recombinant state cannot produce megalomicea. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14153 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, there gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108 51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megM forward PCR primer SEQ ID NO:7.

L3 ANSWER 22 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, there gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

AN ADI14162 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant

nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying enzyme chosen from MegR, **MegF**, **MegK**, **MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, **MegL**, and **MegM** enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14162 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: M. megalomicea megDIV reverse PCR primer SEQ ID NO:16.

L3 ANSWER 23 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
AN ADI14152 DNA DGENE  
AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying enzyme chosen from MegR, **MegF**, **MegK**, **MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, **MegL**, and **MegM** enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14152 DNA DGENE  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying enzyme e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megL reverse PCR primer SEQ ID NO:6.

L3 ANSWER 24 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
**polyketide modifying gene**, where gene encodes  
polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM  
enzymes useful for producing modified polyketide.

AN ADI14164 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying **enzyme** chosen from MegR, **MegF**, **MegK**,  
**MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, MegL, and  
MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence represents a PCR primer used to amplify a gene of the invention.

ACCESSION NUMBER: ADI14164 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.

INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H

PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.

PATENT INFO: WO 2004003169 A2 20040108

51

APPLICATION INFO: WO 2003-US20681 20030630

PRIORITY INFO: US 2002-393016P 20020628

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 2004-203379 [19]

DESCRIPTION: M. megalomicea megBV reverse PCR primer SEQ ID NO:18.

L3 ANSWER 25 OF 27 DGENE COPYRIGHT 2006 The Thomson Corp on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
**polyketide modifying gene**, where gene encodes  
polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM  
enzymes useful for producing modified polyketide.

AN ADI14150 DNA DGENE

AB The invention relates to a novel isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying **enzyme** chosen from MegR, **MegF**, **MegK**,  
**MegCIV**, **MegCV**, **MegBVI**, **MegBIII**, MegL, and  
MegM enzymes. A method of the invention is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising the recombinant nucleic acid under conditions in which the cell expresses a product of a gene encoded by the nucleic acid under conditions in which the unmodified polyketide is present, and producing the modified polyketide. The cell produces megosamine and can attach megosamine to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megosamine. The present sequence is used in the exemplification of the invention.

ACCESSION NUMBER: ADI14150 DNA DGENE

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, where gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes

useful for producing modified polyketide.  
INVENTOR: Hutchinson R C; Katz L; Reid R; Hu Z; Gramajo H  
PATENT ASSIGNEE: (KOSA-N) KOSAN BIOSCIENCES INC.  
PATENT INFO: WO 2004003169 A2 20040108 51  
APPLICATION INFO: WO 2003-US20681 20030630  
PRIORITY INFO: US 2002-393016P 20020628  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
OTHER SOURCE: 2004-203379 [19]  
DESCRIPTION: Synthetic oligonucleotide of the invention SEQ ID NO:4.

L3 ANSWER 26 OF 27 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
**polyketide modifying gene**, where gene encodes  
polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM  
enzymes useful for producing modified polyketide.  
AN 2004-203379 [19] WPIDS  
AB WO2004003169 A UPAB: 20040318  
NOVELTY - An isolated, purified, or recombinant nucleic acid (I)  
comprising a **polyketide modifying gene**,  
where the gene encodes a polyketide modifying **enzyme** chosen from  
MegR, **MegF**, **MegK**, **MegCIV**, **MegCV**,  
MegBVI, **MegBIII**, MegL, and MegM enzymes, is new.  
DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the  
following:  
(1) an isolated, purified, or recombinant nucleic acid (II)  
comprising genes for the biosynthesis mycarose for attachment to a  
polyketide, the enzymes comprising the MegM, MegL, **MegBIII**,  
MegBIV, MegDIV, MegBII-2, and MegBVI enzymes;  
(2) an isolated, purified, or recombinant nucleic acid (III)  
comprising genes for the biosynthesis mycarose for attachment of  
megasamine of a polyketide, the enzymes comprising the MegM, MegL, MegCII,  
MegBVI, MegDIV, MegDV, MegDII, MegDIII, and MegDI enzymes;  
(3) an isolated, purified, or recombinant nucleic acid (IV)  
comprising genes for the biosynthesis of desosamine to a polyketide, the  
enzymes consisting of the MegM, MegL, MegCII, **MegCIV**,  
**MegCV**, MegDII, and MegDIII enzymes;  
(4) an expression vector (V) comprising (I);  
(5) a host cell comprising (I);  
(6) a host cell comprising (II) that expresses a polyketide modifying  
**enzyme** encoded by a gene from a mycarose biosynthetic gene set,  
where the **enzyme** is chosen from MegM, MegL, **MegBIII**,  
MegBIV, MegDIV, MegBII-2, and MegBVI, MegBV, and **MegF**;  
(7) a host cell comprising (III) that expresses a polyketide  
modifying **enzyme** encoded by a gene from a megasamine  
biosynthetic gene set, where the **enzyme** is chosen from MegM,  
MegL, MegCII, MegBVI, MegDIV, MegDV, MegDVI, MegDVII, MegDII, MegDIII, and  
MegDI; and  
(8) a host cell comprising (IV) that expresses a polyketide modifying  
**enzyme** encoded by a gene from a desosamine biosynthetic gene set,  
where the **enzyme** is chosen from MegM, MegL, MegCII,  
**MegCIV**, **MegCV**, MegDII, and MegDIII, and MegCIII.

USE - (M1) is useful for producing a modified polyketide, which  
involves culturing a recombinant cell comprising (I) under conditions in  
which the cell expresses a product of a gene encoded by (I) under  
conditions in which the unmodified polyketide is present, and producing  
the modified polyketide. In (M1), the cell further comprises (I) one or  
more module of a polyketide synthase. The cell produces megasamine and can  
attach megasamine to a polyketide, where the cell, it its naturally  
occurring non-recombinant state cannot produce megasamine. (All claimed.)

DESCRIPTION OF DRAWING(S) - The drawing shows a schematic of the  
megalomicin polyketide synthase (meg DEBS) and corresponding meg genes  
upstream and downstream of the meg DEBS region and cosmids overlapping

this region.

Dwq. 1/3

ACCESSION NUMBER: 2004-203379 [19] WPIDS  
DOC. NO. CPI: C2004-080057  
TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, there gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide.  
DERWENT CLASS: B03 B04 C02 D16  
INVENTOR(S): GRAMAJO, H; HU, Z; HUTCHINSON, R C; KATZ, L; REID, R; HUTCHINSON, C R  
PATENT ASSIGNEE(S): (KOSA-N) KOSAN BIOSCIENCES INC; (GRAM-I) GRAMAJO H; (HUZZ-I) HU Z; (HUTC-I) HUTCHINSON C R; (KATZ-I) KATZ L; (REID-I) REID R  
COUNTRY COUNT: 105  
PATENT INFORMATION:

PATENT NO	KIND	DATE	WEEK	LA	PG
WO 2004003169	A2	20040108	(200419)*	EN	51
RW:	AT BE BG CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE LS LU MC MW MZ NL OA PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW				
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AU 2003258978	A1	20040119	(200447)		
US 2004203015	A1	20041014	(200468)		

**APPLICATION DETAILS:**

PATENT NO	KIND	APPLICATION	DATE
WO 2004003169	A2	WO 2003-US20681	20030630
AU 2003258978	A1	AU 2003-258978	20030630
US 2004203015	A1 Provisional	US 2002-393016P US 2003-611442	20020628 20030630

**FILING DETAILS:**

PATENT NO	KIND	PATENT NO
AU 2003258978	A1 Based on	WO 2004003169

PRIORITY APPLN. INFO: US 2002-393016P 20020628; US  
2003-611442 20030630

L3 ANSWER 27 OF 27 BIOTECHDS COPYRIGHT 2006 THE THOMSON CORP. on STN  
TI Novel isolated, purified, or recombinant nucleic acid comprising  
**polyketide modifying gene**, there gene encodes  
polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM  
enzymes useful for producing modified polyketide;  
involving vector-mediated gene transfer and expression in host cell  
for polyketide production

AN 2004-10434 BIOTECHDS

**AB DERWENT ABSTRACT:**

NOVELTY - An isolated, purified, or recombinant nucleic acid (I) comprising a **polyketide modifying gene**, where the gene encodes a polyketide modifying **enzyme** chosen from MegR, MegF, MegK, MegCIV, MegCV, MegBVI, MegBIII, MegL, and MegM enzymes, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following: (1) an isolated, purified, or recombinant nucleic acid (II) comprising genes for the biosynthesis mycarose for attachment to a polyketide, the enzymes comprising the MegM, MegL, **MegBIII**, MegBIV, MegDIV, MegBII-2, and MegBVI enzymes; (2) an isolated, purified, or recombinant nucleic acid (III) comprising genes for the biosynthesis mycarose for attachment of megosamine of a polyketide, the enzymes comprising the MegM, MegL, MegCII, MegBVI, MegDIV, MegDV, MegDII, MegDIII, and MegDI enzymes; (3) an isolated, purified, or recombinant nucleic acid (IV) comprising genes for the biosynthesis of desosamine to a polyketide, the enzymes consisting of the MegM, MegL, MegCII, **MegCIV**, **MegCV**, MegDII, and MegDIII enzymes; (4) an expression vector (V) comprising (I); (5) a host cell comprising (I); (6) a host cell comprising (II) that expresses a polyketide modifying **enzyme** encoded by a gene from a mycarose biosynthetic gene set, where the **enzyme** is chosen from MegM, MegL, **MegBIII**, MegBIV, MegDIV, MegBII-2, and MegBVI, MegBV, and **MegF**; (7) a host cell comprising (III) that expresses a polyketide modifying **enzyme** encoded by a gene from a megalomycin biosynthetic gene set, where the **enzyme** is chosen from MegM, MegL, MegCII, MegBVI, MegDIV, MegDV, MegDVI, MegDVII, MegDII, MegDIII, and MegDI; and (8) a host cell comprising (IV) that expresses a polyketide modifying **enzyme** encoded by a gene from a desosamine biosynthetic gene set, where the **enzyme** is chosen from MegM, MegL, MegCII, **MegCIV**, **MegCV**, MegDII, and MegCIII.

BIOTECHNOLOGY - Preferred Nucleic Acid: In (I), the gene encodes a polyketide modifying enzymes chosen from MegR, **MegK**, **MegCV**, **MegCIV**, MegBVI, **MegF**, MegBII, MegM, and MegL. (I) further comprises gene encoding an **enzyme** for the attachment of mycarose to the polyketide, preferably MegBV **enzyme**. (I) further comprises gene encoding an **enzyme** for hydroxylation of the polyketide, preferably **MegF enzyme**. (IV) further comprises gene encoding an **enzyme** for the attachment of desosamine to the polyketide, preferably MegCIII **enzyme**. The **Polyketide modifying gene** is operably linked to heterologous promoter.

USE - (M1) is useful for producing a modified polyketide, which involves culturing a recombinant cell comprising (I) under conditions in which the cell expresses a product of a gene encoded by (I) under conditions in which the unmodified polyketide is present, and producing the modified polyketide. In (M1), the cell further comprises (I) one or more module of a polyketide synthase. The cell produces megalomycin and can attach megalomycin to a polyketide, where the cell, in its naturally occurring non-recombinant state cannot produce megalomycin. (All claimed.)

EXAMPLE - Isolation of the megalomycin biosynthetic gene cluster was as follows. A cosmid library was prepared in SuperCos vectors from Micromonospora megalomicina total DNA partially digested with Sau3AI and introduced into Escherichia coli using a Gigapack III XL in vitro packaging kit. 32P-labeled DNA probes encompassing the KS2 domain from DEBS, or a mixture of segments encompassing modules 1 and 2 from DEBS, were used separately to screen the cosmid library by colony hybridization. Several colonies which hybridized with the probes were further analyzed by sequencing the ends of their cosmid inserts using T3 and T7 primers. BLAST analysis of the sequences revealed several colonies with DNA sequences highly homologous to genes from the ery cluster. Together with restriction analysis, that led to the isolation of two overlapping cosmids, pKOSO79-93A and pKOSO79-93D which covered 45 kb of the meg cluster. A 400 base pair PCR fragment was generated from the left end of pKOSO79-93D and used to reprobe the cosmid library. Likewise, a 200 base pair PCR fragment generated from the right end of pKOSO79-93A was used to reprobe the cosmid library. Analysis of hybridizing colonies, as described above, resulted in identification of two additional cosmids pKOSO79-93B adjacent to the 5' end of pKOSO79-93D and pKOSO05.57-2.3B.

which overlapped the 3' ends of pKOSO79-93A and pKOSO79-93D cosmids. BLAST analysis of the far left and right end sequences of these cosmids indicated no homology to any known genes related to polyketide biosynthesis, and therefore indicates that the set of four cosmids spans the entire megalomicin biosynthetic gene cluster. The glycosyl synthase, transfer, and regulatory genes of the upstream region of the meg PKS were contained in the nucleotide sequence which had a 9024 nucleotide sequence given in the specification. The glycosyl synthase, transfer, and regulatory genes of the downstream region of the meg PKS were contained in the nucleotide sequence which had a 17596 nucleotide sequence given in the specification. (51 pages)

ACCESSION NUMBER: 2004-10434 BIOTECHDS

TITLE: Novel isolated, purified, or recombinant nucleic acid comprising **polyketide modifying gene**, there gene encodes polyketide modifying **enzyme** e.g., MegR, **MegK**, or MegM enzymes useful for producing modified polyketide; involving vector-mediated gene transfer and expression in host cell for polyketide production

AUTHOR: HUTCHINSON R C; KATZ L; REID R; HU Z; GRAMAO H

PATENT ASSIGNEE: KOSAN BIOSCIENCES INC

PATENT INFO: WO 2004003169 8 Jan 2004

APPLICATION INFO: WO 2003-US20681 30 Jun 2003

PRIORITY INFO: US 2002-393016 28 Jun 2002; US 2002-393016 28 Jun 2002

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: WPI: 2004-203379 [19]

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E2	1	HUTCHINSON Z/AU
E3	0 -->	HUTCHINSON, C/AU
E4	1	HUTCHINSONCOLAS J/AU
E5	1	HUTCHINSONCOLE H/AU
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=> e katz, l/au

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